What is claimed is:

1. A method of treating cancer comprising administering to a patient in need thereof an effective amount of a compound of Formula I:

wherein,

each R is independently hydrogen, hydroxy, alkyl, aryl, cycloalkyl, heteroaryl, alkoxy, heterocyclic or amino;

each R_1 is independently alkyl, halo, alkoxy, haloalkyl, haloalkoxy, cycloalkyl, heterocyclic, hydroxy, -C(O)- R_8 , -NR₉R₁₀, -NR₉C(O)- R_{12} or -C(O)NR₉R₁₀;

each R_2 is independently alkyl, aryl, heteroaryl, $-C(O)-R_8$ or SO_2R ", where R" is alkyl, aryl, heteroaryl, NR_9N_{10} or alkoxy;

each R_5 is independently hydrogen, alkyl, aryl, haloalkyl, cycloalkyl, heteroaryl, heterocyclic, hydroxy, -C(O)- R_8 or (CHR)_r R_{11} ;

X is O or S; j is 0 or 1; p is 0, 1, 2 or 3; q is 0, 1 or 2; r is 0, 1, 2 or 3;

R₈ is hydroxy, alkyl, aryl, heteroaryl, alkoxy, cycloalkyl or heterocyclic;

 R_9 and R_{10} are independently hydrogen, alkyl, aryl, aminoalkyl, heteroaryl, cycloalkyl and heterocyclic, or R_9 and R_{10} together with N may form a ring, where the ring atoms are selected from the group consisting of C, N, O and S;

R₁₁ is hydroxy, amino, monosubstituted amino, disubstituted amino, alkyl, aryl, heteroaryl, alkoxy, cycloalkyl or heterocyclic

R₁₂ is alkyl, aryl, heteroaryl, alkoxy, cycloalkyl or heterocyclic; and

Z is hydroxy, -O-alkyl, or $-NR_3R_4$, where R_3 and R_4 are independently hydrogen, alkyl, aryl, heteroaryl, cycloalkyl, or heterocyclic, or R_3 and R_4 may combine with N to form a ring where the ring atoms are selected from the group consisting of CH_2 , N, O and S, or

wherein Y is independently CH₂, O, N or S, Q is C or N, n is independently 0, 1, 2, 3 or 4, and m is 0, 1, 2 or 3;

or a pharmaceutically acceptable salt, hydrate or solvate thereof, in combination with at least one chemotherapeutic agent selected from the group consisting of microtubule interference agents, topoisomerase inhibitors, alkylating agents, thymidylate synthase inhibitors, irreversible steroidal aromatase inactivators, anti-metabolites, pyrimidine antagonists, purine antagonists, ribonucleotide reductase inhibitors, and kinase inhibitors.

- 2. The method of claim 1, wherein R_1 is halo and p is 1.
- 3. The method of claim 1, wherein R_1 is F or Cl and p is 1.
- 4. The method of claim 1, wherein Z is $-NR_3R_4$ wherein R_3 and R_4 are lower alkyl or form a morpholine ring.

5. The method of claim 1, wherein Z is:

$$---N \xrightarrow{(Y)_n} Q \xrightarrow{\begin{pmatrix} R^1 \\ C \\ R^1 \end{pmatrix}_m} R^3$$

wherein each Y is CH₂, each n is 2, m is 0 and R₃ and R₄ form a morpholine ring.

- 6. The method of claim 1, wherein R_2 is methyl and q is 2, wherein the methyls are bonded at the 3 and 5 positions.
- 7. The method of claim 1, wherein the compound of formula I is selected from the group consisting of

and pharmaceutically acceptable salts, solvates and hydrates thereof.

8. The method of claim 1, wherein the compound of formula I is selected from the group consisting of:

and pharmaceutically acceptable salts, solvates and hydrates thereof.

9. The method of claim 1, wherein the compound of Formula (I) is:

or a pharmaceutically acceptable salt, solvate or hydrate thereof.

- 10. The method of claim 9, wherein the salt is a malate salt.
- 11. The method of claim 1, wherein the at least one chemotherapeutic agent is selected from the group consisting of taxanes, vinca alkyloids, topoisomerase I inhibitors and topoisomerase II inhibitors.
- 12. The method of claim 1, wherein the at least one chemotherapeutic agent is selected from the group consisting of paclitaxel, docetaxel, vinblastine, vincristine, vindesine, irinotecan, doxorubicin, epirubicin, leucovorin, etopside, teniposide, idarubicine, gemcitabine, daunorubicin, carboplatin, cisplatin, oxaliplatin, chlorambucil, melphalan, cyclophosphamide, ifosfamide, temozolomide, thiotepa, mitomycin C, busulfan, carmustine, lomustine, 5-fluorouracil, capecitabine, exemestane, methotrexate, trimetrexate, fluorouracil, fluorodeoxyuridine, azacytidine, mercaptopurine, thioguanine, pentostatin, cytarabine, fludarabine, hydroxyurea, bevacizumab, cetuximab, gefitinib and imatinib.
- 13. The method of claim 1, wherein the cancer is breast cancer, small cell lung carcinoma, colon cancer, non-small cell lung cancer, renal cell cancer, a gastrointestinal stromal tumor, thyroid cancer, a sarcoma or a neuroendocrine tumor.

- 14. The method of claim 1, wherein the cancer is non-small cell lung cancer and the at least one chemotherapeutic agent is carboplatin and paclitaxel.
- 15. The method of claim 1, wherein the cancer is non-small cell lung cancer and the at least one chemotherapeutic agent is carboplatin, taxotere, cisplatin, gemcitabine, 5-fluorouracil, irinotecan or leucovorin.
- 16. The method of claim 1, wherein the cancer is colon cancer and the at least one chemotherapeutic agent is 5-fluorouracil, oxaliplatin or leucovorin.
- 17. A method of treating cancer comprising administering to a patient in need thereof an effective amount of a compound selected from the group consisting of:
- 5-(5-Fluoro-2-oxo-1,2-dihydro-indol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylamino-ethyl)-amide;
- 5-(5-Fluoro-2-oxo-1,2-dihydro-indol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-pyrrolidin-1-yl-ethyl)-amide;
- 5-(5-Fluoro-2-oxo-1,2-dihydro-indol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-morpholin-4-yl-ethyl)-amide;
- (S)-5-(5-Fluoro-2-oxo-1,2-dihydro-indol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-hydroxy-3-morpholin-4-yl-propyl)-amide;
- (R)-5-(5-Fluoro-2-oxo-1,2-dihydro-indol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-hydroxy-3-morpholin-4-yl-propyl)-amide;
- 5-(5-Fluoro-2-oxo-1,2-dihydro-indol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-hydroxy-3-morpholin-4-yl-propyl)-amide;
- 5-(5-Chloro-2-oxo-1,2-dihydro-indol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-hydroxy-3-morpholin-4-yl-propyl)-amide;
- 5-(5-Fluoro-2-oxo-1,2-dihydro-indol-3-ylidenemethyl)-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-ethylamino-ethyl)-amide; and
- 3-[3,5-dimethyl-4-(4-morpholin-4-yl-piperidine-1-carbonyl)-1H-pyrrol-2-methylene]-5-fluoro-1,3-dihydro-indol-2-one,

- or a pharmaceutically acceptable salt, hydrate or solvate thereof, in combination with at least one chemotherapeutic agent selected from the group consisting of microtubule interference agents, topoisomerase inhibitors, alkylating agents, thymidylate synthase inhibitors, irreversible steroidal aromatase inactivators, anti-metabolites, pyrimidine antagonists, purine antagonists, ribonucleotide reductase inhibitors, and kinase inhibitors.
- 18. The method of claim 17, wherein the at least one chemotherapeutic agent is selected from the group consisting of taxanes, vinca alkyloids, topoisomerase I inhibitors and topoisomerase II inhibitors.
- 19. The method of claim 17, wherein the at least one chemotherapeutic agent is selected from the group consisting of paclitaxel, docetaxel, vinblastine, vincristine, vindesine, irinotecan, doxorubicin, epirubicin, leucovorin, etopside, teniposide, idarubicine, gemcitabine, daunorubicin, carboplatin, cisplatin, oxaliplatin, chlorambucil, melphalan, cyclophosphamide, ifosfamide, temozolomide, thiotepa, mitomycin C, busulfan, carmustine, lomustine, 5-fluorouracil, capecitabine, exemestane, methotrexate, trimetrexate, fluorouracil, fluorodeoxyuridine, azacytidine, mercaptopurine, thioguanine, pentostatin, cytarabine, fludarabine, hydroxyurea, bevacizumab, cetuximab, gefitinib and imatinib.
- 20. The method of claim 17, wherein the cancer is breast cancer, small cell lung carcinoma, colon cancer, non-small cell lung cancer, renal cell cancer, a gastrointestinal stromal tumor, thyroid cancer, a sarcoma or a neuroendocrine tumor.